

This listing of claims will replace all prior versions, and listings, of claims in the application:

**LISTING OF CLAIMS:**

**1. - 24. (Canceled)**

**25. (Currently Amended)** A composition comprising:

i) a granulated preparation comprising a complex between an estrogen and a cyclodextrin;  
and

ii) optionally, one or more excipients,

the composition having a stability such that said estrogen is in an amount of at least ~~90%~~ 85% w/w in relation to the initial content of said estrogen after storage for 12 months at 40°C and 75% relative humidity (RH); and

the composition being essentially free of polyvinylpyrrolidone.

**26. – 27. (Canceled)**

**28. (Original)** The composition according to claim 25, wherein the estrogen is in an amount from about 0.002 % w/w to 2% w/w.

**29. (Original)** The composition according to claim 25, wherein the estrogen is in an amount from about 0.004% w/w to 0.2% w/w.

**30. (Original)** The composition according to claim 25, wherein the estrogen is ethinyl estradiol and the cyclodextrin is  $\beta$ -cyclodextrin, the ethinyl estradiol is in an amount relative to the ethinyl estradiol- $\beta$ -cyclodextrin complex of from about 5 % w/w to 20 % w/w.

**31. (Original)** The composition according to claim 25, wherein the estrogen is ethinyl estradiol and the cyclodextrin is  $\beta$ -cyclodextrin, the ethinyl estradiol is in an amount relative to the ethinyl estradiol- $\beta$ -cyclodextrin complex of from about 8% w/w to 15% w/w.

**32. - 36. (Canceled)**

**37. (Currently Amended)** A method for improving the stability of an estrogen in a pharmaceutical composition that comprises an estrogen and one or more excipients in a granulate preparation, which method comprises:

- i) forming a complex between said estrogen and a cyclodextrin; and
- ii) granulating providing the complex in a granulator to provide the granulate preparation;
- iii) ~~introducing a granulating liquid into the granulator; and~~
- iv) ~~mixing the granulator contents under granulation conditions with the one or more excipients to provide a granulate, such that the relative humidity of the granulate does not exceed 60%, as determined at a temperature between 20°C and 40°C,~~

provided that, the composition is prepared so that it is essentially free of polyvinylpyrrolidone ~~when the one or more excipients comprises polyvinylpyrrolidone, it is present in an amount of at most 2% w/w.~~

**38. - 41. (Canceled)**

**42. (Currently Amended)** A process for manufacturing a granulate preparation comprising a complex between an estrogen and a cyclodextrin, which comprises:

- i) loading the complex, optionally ~~further~~ one or more other therapeutically active agent(s), and one or more excipients into a granulator;
- ii) applying a granulating liquid onto the loaded complex and the one or more excipients, and
- iii) granulating and drying, ~~under granulation conditions~~ so as to obtain granules having a relative humidity not exceeding 60%, as determined at a temperature between 20°C and 40°C.

**43. (Original)** The process according to claim 42, wherein the complex and the optionally further one or more therapeutically active agent(s) are provided as individual agent(s) without being pre-mixed with excipients.

**44. (Original)** The process according to claim 42, wherein the relative humidity of the granulate preparation does not exceed 55%, as determined at a temperature between 20°C and 40°C.

**45. (Original)** The process according to claim 42, wherein the relative humidity of the granulate preparation does not exceed 45%, as determined at a temperature between 20°C and 40°C.

**46. (Original)** The process according to claim 42, wherein the relative humidity of the granulate preparation does not exceed 40%, as determined at a temperature between 20°C and 40°C.

**47. (Canceled)**

**48. (Previously presented)** The composition according to claim 25, wherein the composition comprises one or more excipient(s) which is/are selected from the group consisting of starch, cellulose, hydroxypropylcellulose, hydroxypropylmethylcellulose and maltodextrin.

**49. (Previously presented)** The composition according to claim 25, wherein the estrogen is selected from the group consisting of ethinyl estradiol, estradiol, estradiol sulfamates, estradiol valerate, estradiol benzoate, estrone, estrone sulfate and mixtures thereof.

**50. (Previously presented)** The composition according to claim 49, wherein the estrogen is ethinyl estradiol.

**51. (Previously presented)** The composition according to claim 25, wherein the cyclodextrin is selected from the group consisting of  $\alpha$ -cyclodextrin,  $\beta$ -cyclodextrin,  $\gamma$ -cyclodextrin and alkylated or acylated derivatives thereof.

- 52. (Previously presented)** The composition according to claim 25, wherein the cyclodextrin is  $\beta$ -cyclodextrin or an alkylated or acylated derivative thereof.
- 53. (Previously presented)** The composition according to claim 25, wherein the estrogen is in an amount relative to the cyclodextrin such that a molar ratio between the estrogen and the cyclodextrin is from about 2:1 to 1:10.
- 54. (Currently Amended)** The composition according to claim 25, further comprising one or more other therapeutically active agent(s).
- 55. (Currently Amended)** The composition according to claim 54, wherein the one or more other therapeutically active agent(s) is a progestogen.
- 56. (Currently Amended)** The composition according to claim ~~55~~ 56, wherein the progestogen is selected from the group consisting of drospirenone, levonorgestrel, norgestrel, gestodene, dienogest, cyproterone acetate, norethisterone, norethisterone acetate, desorgestrel, and 3-keto-desorgestrel.
- 57. (Previously presented)** The composition according to claim 56, wherein the progestogen is drospirenone.
- 58. (Previously presented)** The composition according to claim 57, wherein drospirenone is in micronized form.
- 59. (Previously presented)** The composition according to claim 58, wherein drospirenone is in an amount from about 0.4 % to 20% w/w.
- 60. (Previously presented)** The composition according to claim 25, wherein the complex is micronized.
- 61. (Previously presented)** The composition according to claim 25, further comprising an antioxidant.

- 62. (Previously presented)** The composition of claim 25, wherein the granulated preparation has a mean particle size of at least about 100  $\mu\text{m}$ .
- 63. (Previously presented)** The method according to claim 37, provided that the one or more excipient(s) is/are selected from the group consisting of starch, cellulose, hydroxypropylcellulose, hydroxypropylmethylcellulose and maltodextrin.
- 64. (Previously presented)** The process according to claim 42, provided that, when the granulated preparation comprises polyvinylpyrrolidone, it is present in an amount of at most 2% w/w.
- 65. (Previously presented)** The method according to claim 42, provided that the one or more excipient(s) is/are selected from the group consisting of starch, cellulose, hydroxypropylcellulose, hydroxypropylmethylcellulose and maltodextrin.
- 66. (Previously presented)** The process according to claim 42, wherein the estrogen is selected from the group consisting of ethinyl estradiol (EE), estradiol, estradiol sulfamates, estradiol valerate, estradiol benzoate, estrone, estrone sulfate and mixtures thereof.
- 67. (Previously presented)** The process according to claim 66, wherein the estrogen is ethinyl estradiol.
- 68. (New)** The process of claim 42, wherein the granulating is by fluidized bed granulation.